- 1. A method for diagnosing a malignant neoplasm in a mammal, comprising contacting a bodily fluid from said mammal with an antibody which binds to an human aspartyl (asparaginyl) beta-hydroxylase (HAAH) polypeptide under conditions sufficient to form an antigen-antibody complex and detecting the antigen-antibody complex.
- 2. The method of claim 1, wherein said neoplasm is derived from endodermal tissue.

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- 3. The method of claim 1, wherein said neoplasm is selected from the group consisting of colon cancer, breast cancer, pancreatic cancer liver cancer, and cancer of the bile ducts.
- 4. The method of claim 1, wherein said neoplasm is a cancer of the central nervous system (CNS).
- 5. The method of claim 1, wherein said bodily fluid is selected from the group consisting of a CNS-derived bodily fluid, blood, serum, urine, saliva, sputum, lung effusion, and ascites fluid.
- 1 6. The method of claim 1, wherein said antibody is a 2 monoclonal antibody.
- 7. The method of claim 6, wherein said monoclonal antibody is FB50.
- 8. The method of claim 6, wherein said monoclonal antibody is selected from the group consisting of 5C7, 5E9, 3 19B, 48A, 74A, 78A, 86A.

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- A method for prognosis of a malignant neoplasm 1 2 of a mammal, comprising
- (a) contacting a bodily fluid from said mammal 3 4 with an antibody which binds to an HAAH polypeptide under conditions sufficient to form an antigen-antibody complex 5 and detecting the antigen-antibody complex; 6
- 7 (b) quantitating the amount of complex to determine the level of HAAH in said fluid; and
- (c) comparing the level of HAAH in said fluid 9 with a normal control level of HAAH, wherein increasing 10 levels of HAAH over t_{a} me indicates an adverse prognosis. 11
 - 10. A method of $\$ inhibiting tumor growth in a mammal comprising administering to said mammal a compound which inhibits expression of HAAH.
 - 11. The method of claim 10, wherein said compound is a HAAH antisense nucleic ac
 - The method of claim 10, wherein said compound 12. is a ribozyme.
 - The method of claim 10, wherein said tumor is 1 derived from endodermal tissue. 2
 - The method of claim 1d, wherein said tumor is 1 selected from the group consisting of colon cancer, breast 2 cancer, pancreatic cancer, liver cancer, and cancer of the bile ducts. 4
 - The method of claim 10, wherein said tumor is a 15. 1 2 CNS tumor.

- 1 16. A method of inhibiting tumor growth in a mammal comprising administering to said mammal a compound which inhibits an enzymatic activity of HAAH.
- 1 17. The method of claim 16, wherein said enzymatic 2 activity is hydroxylase activity...
- 1 18. The method of claim 16, wherein said compound 2 is a dominant negative mutant of HAAH.
- 1 19. The method of claim 18, wherein said dominant 2 negative mutant HAAH comprises a mutation in a catalytic 3 domain of HAAH.

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- 20. The method of claim 16, wherein said compound is an HAAH-specific introopdy.
- 21. The method of claim 16, wherein said compound is L-mimosine.
- 22. The method of claim 16, wherein said compound is a hydroxypyridone.
- 1 23. A method of inhibiting tumor growth in a mammal 2 comprising administering to said mammal a compound which 3 inhibits signal transduction through the IRS signal 4 transduction pathway.
 - 1 24. The method of claim 23, wherein said compound 2 inhibits IRS phosphorylation.
 - 1 25. The method of claim 23 wherein said compound 2 inhibits binding of Fos or Jun to an HAAH promoter sequence.

- 26. A method of inhibiting tumor growth in a mammal comprising administering to said mammal a compound which inhibits HAAH hydroxylation of a NOTCH polypeptide.
- 27. The method of claim 26, wherein said compound inhibits hydroxylation of an EGF like repeat sequence in a NOTCH polypeptide.
- 28. A method of killing a tumor cell comprising
 contacting said tumor cell with cytotoxic agent linked to an
 HAAH-specific antibody.
 - 29. A monoclonal antibody that binds to an epitope of HAAH.
 - 30. The antibody of claim 29, wherein said epitope is within a catalytic site of $\widehat{\text{HAAH}}$.

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- 31. The antibody of claim 29, wherein said monoclonal antibody is selected from the group consisting of 5C7, 5E9, 19B, 48A, 74A, 78A, 86A.
- 1 32. The antibody of claim 29, wherein said 2 monoclonal antibody is selected from the group consisting of 3 HA238A, HA221, HA239, HA241, HA329, or HA355.
 - 33. A composition comprising a monoclonal antibody that binds to an epitope of HAAH linked to a cytotoxic agent, wherein said composition preferentially kills tumor cells compared to non-tumor cells.
 - 34. A kit for diagnosis of a tumor in a mammal, comprising the antibody of claim 29.

The kit of claim 34, wherein said antibody is 2 immobilized on a solid phase. 1 The kit of claim 35, wherein said solid phase is selected from a group consisting of an assay plate, an 2 assay well, a nitrocellulose membrane, a bead, a dipstick, 3 4 and a component of an elution column. \setminus A method of determining whether a candidate 1 2 compound inhibits HAAH enzymatic activity, comprising 3 (a) providing a HAAH polypeptide; (b) providing a polypeptide comprising an EGF-like 4 5 domain; (c) contacting said HAAH polypeptide or said NOTCH polypeptide with\said candidate compound; (d) determining hydroxylation of said polypeptide of step (b), wherein a decrease in hydroxylation in the presence of said candidate compound compared to that in the # **11** absence of said compound indicates that said compound 12 inhibits HAAH enzymatic activity. ı.i ,± 1 38. A method of determining whether a candidate compound inhibits HAAH activation of NOTCH, comprising <u>...</u> 2 (a) providing a dell expressing HAAH; 3 (b) contacting said cell with a candidate compound; 4 5 and (c) measuring trans \(\)ocation of activated NOTCH to 6

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translocation in the presence of said compound compared to

that in the absence of said compound indicates that said

the nucleus of said cell, wherein a decrease in

compound HAAH activation of NOTCH.